

Attorney's Docket No.: 07039-281002

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Christopher B. Eckman et al.      Art Unit : Unknown  
Serial No. : 10/681,609      Examiner : Unknown  
Filed : October 8, 2003  
Title : METHODS OF REDUCING BETA-AMYLOID POLYPEPTIDES

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

TRANSMITTAL

The following correspondence relating to this application is enclosed for filing:

1. Information Disclosure Statement (1 page);
2. Form PTO-1449 (3 pages);
3. Copies of references (10); and
3. A return postcard.

Please date stamp and mail the enclosed postcard.

Please apply any charges or credits to Deposit Account No. 06-1050.

Respectfully submitted,

Date: March 16, 2004

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INFORMATION DISCLOSURE STATEMENT

Applicants submit the references listed on the attached Form PTO-1449.

Under 35 USC §120, this application relies on the earlier filing date of U.S. Application No/ 09/804,420, filed on March 12, 2001. Only the references that were not submitted to and/or cited by the Office in the prior application are enclosed.

This statement is being filed before the receipt of a first Office Action on the merits.

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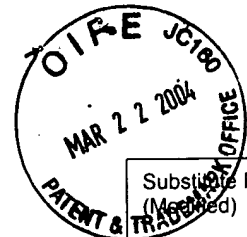
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Substitute Form PTO-1449 (Modified)	U.S. Department of Commerce Patent and Trademark Office	Attorney's Docket No. 07039-281002	Application No. 10/681,609
<b>Information Disclosure Statement by Applicant</b> (Use several sheets if necessary) (37 CFR §1.98(b))		Applicant Christopher B. Eckman et al.	
		Filing Date October 8, 2003	Group Art Unit

U.S. Patent Documents							
Examiner Initial	Desig. ID	Document Number	Publication Date	Patentee	Class	Subclass	Filing Date If Appropriate
	AA	4,666,829	05/19/87	Glenner et al.			
	AB	5,538,845	07/23/96	Knops et al.			
	AC	5,552,426	09/03/96	Lunn et al.			
	AD	5,604,102	02/18/97	McConlogue et al.			
	AE	5,703,129	12/30/97	Felsenstein et al.			
	AF	5,766,846	06/16/98	Schlossmacher et al.			
	AG	6,096,782	08/01/00	Audia et al.			
	AH	6,191,166	02/20/01	Audia et al.			
	AI	6,211,235	04/03/01	Wu et al.			

Foreign Patent Documents or Published Foreign Patent Applications								
Examiner Initial	Desig. ID	Document Number	Publication Date	Country or Patent Office	Class	Subclass	Translation	
							Yes	No
	AJ	JP59020298A	02/01/84	Japan				
	AK	KR9300059B	01/08/93	Korea				

Other Documents (include Author, Title, Date, and Place of Publication)		
Examiner Initial	Desig. ID	Document
	AL	Asami-Odaka et al., "Long Amyloid $\beta$ -Protein Secreted from Wild-Type Human Neuroblastoma IMR-32 Cells," <u>Biochem.</u> , 1995, 34:10272-10278
	AM	Brinker, "Variations in Effective Botanical Products – The Case for Diversity of Forms for Herbal Preparations as Supported by Scientific Studies," <u>HerbalGram</u> , 1999, 46:36-50
	AN	Buckley, "Separation Techniques," School of Education and Humanities, North East Wales Institute of Higher Education at <a href="http://www.newi.ac.uk/buckley/separ.htm">http://www.newi.ac.uk/buckley/separ.htm</a>
	AO	Buxbaum et al., "Processing of Alzheimer $\beta$ /A4 amyloid precursor protein: Modulation by agents that regulate protein phosphorylation," <u>Proc. Natl. Acad. Sci. USA</u> , 1990, 87:6003-6006
	AP	Buxbaum et al., "Protein phosphorylation inhibits production of Alzheimer amyloid $\beta$ /A4 peptide," <u>Proc. Natl. Acad. Sci. USA</u> , 1993, 90:9195-9198
	AQ	Chevallier et al., "Cathepsin D displays in vitro $\beta$ -secretase-like specificity," <u>Brain Res.</u> , 1997, 750:11-19
	AR	Dovey et al., "Functional gamma-secretase inhibitors reduce beta-amyloid peptide levels in brain," <u>J. Neurochem.</u> , 2001, 76:173-181
	AS	Eckman et al., "Degradation of the Alzheimer's Amyloid $\beta$ Peptide by endothelin-converting Enzyme," <u>J. Biol. Chem.</u> , 2001, 276(27):24540-24548

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	AT	Fassbender et al., "Simvastatin strongly reduces levels of Alzheimer's disease $\beta$ -amyloid peptides A $\beta$ 42 and A $\beta$ 40 <i>in vitro</i> and <i>in vivo</i> ," <u>Proc. Natl. Acad. Sci. USA</u> , 2001, 98(10):5856-5861
	AU	Frears et al., "The role of cholesterol in the biosynthesis of $\beta$ -amyloid," <u>NeuroReport</u> , 1999, 10(8):1699-1705
	AV	Gandy, "Neurohormonal Signaling Pathways and the Regulation of Alzheimer $\beta$ -Amyloid Precursor Metabolism," <u>Trends Endocrinol. Metab.</u> , 1999, 10(7):273-279
	AW	Geriatric Nutritional Review – "Black Cohosh," AgeNet Eldercare Network at <a href="http://www.agenet.com/herbal_supplements/blackcohosh.html">http://www.agenet.com/herbal_supplements/blackcohosh.html</a> .
	AX	Glenbrook Farms, "Herb Facts and Tips, Black Cohosh," at <a href="http://www.glenbrookfarm.com/herbs/blackcoh.htm">http://www.glenbrookfarm.com/herbs/blackcoh.htm</a> .
	AY	Gouras et al., "Testosterone reduces neuronal secretion of Alzheimer's $\beta$ -amyloid peptides," <u>Proc. Natl. Acad. Sci. USA</u> , 2000, 97(3):1202-1205
	AZ	Haass and Selkoe, "Cellular Processing of $\beta$ -Amyloid Precursor Protein and the Genesis of Amyloid $\beta$ -Peptide," <u>Cell</u> , 1993, 75:1039-1042
	AAA	Haass et al., "Normal Cellular Processing of the $\beta$ -Amyloid Precursor Protein Results in the Secretion of the Amyloid $\beta$ Peptide and Related Molecules," <u>Ann. N.Y. Acad. Sci.</u> , 695:109-116
	ABB	Haugabook et al., "Reduction of A $\beta$ accumulation in the Tg2576 animal model of Alzheimer's disease after oral administration of the phosphatidyl-inositol kinase inhibitor wortmannin," <u>FASEB J.</u> , 2001, 15:16-18
	ACC	Holcomb et al., "Accelerated Alzheimer-type phenotype in transgenic mice carrying both mutant <i>amyloid precursor protein</i> and <i>presenilin 1</i> transgenes," <u>Nature Med.</u> , 1998, 4:97-100
	ADD	Hung et al., "Activation of Protein Kinase C Inhibits Cellular Production of the Amyloid $\beta$ -Protein," <u>J. Biol. Chem.</u> , 1993, 268(31):22959-22962
	AEE	Iversen et al., "The toxicity <i>in vitro</i> of $\beta$ -amyloid protein," <u>Biochem. J.</u> , 1995, 311:1-16
	AFF	Jacobsen et al., "The Release of Alzheimer's Disease $\beta$ Amyloid Peptide is Reduced by Phorbol Treatment," <u>J. Biol. Chem.</u> , 1994, 269(11):8376-8382
	AGG	King's Amercian Dispensatory, "Cimicifunga (U.S.P.)-Cimicifuga," at <a href="http://www.ibiblio.org/herbmed/eclectic/kings/cimicifuga.html">http://www.ibiblio.org/herbmed/eclectic/kings/cimicifuga.html</a> .
	AHH	Koo, "Phorbol Esters Affect Multiple Steps in $\beta$ -Amyloid Precursor Protein Trafficking and Amyloid $\beta$ -Protein Production," <u>Mol. Med.</u> , 1997, 3(3):204-211
	AII	Lahiri et al., "The secretion of amyloid $\beta$ -peptides is inhibited in the tacrine-treated human neuroblastoma cells," <u>Mol. Brain Res.</u> , 1998, 62:131-140
	AJJ	Lahiri et al., "Cholinesterase inhibitors, $\beta$ -amyloid precursor protein and amyloid $\beta$ -peptides in Alzheimer's disease," <u>Acta Neurol. Scand.</u> , 2000, Suppl. 176:60-67
	AKK	Lieberman, "Evidence-Based Natural Medicine – A Review of the Effectiveness of <i>Cimicifuga racemosa</i> (Black Cohosh) for the Symptoms of Menopause," <u>J. Women's Health</u> , 1998, 7(5):525-529
	ALL	Lim et al., "Ibuprofen Suppresses Plaque Pathology and Inflammation in a Mouse Model for Alzheimer's Disease," <u>J. Neurosci.</u> , 2000, 20(15):5709-5714
	AMM	Masters et al., "Amyloid plaque core protein in Alzheimer disease and Down syndrome," <u>Proc. Natl. Acad. Sci. USA</u> , 1985, 82:4245-4249
	ANN	Murray, "Indigestion, antacids, achlorydria and H. pylori," <u>Am. J. Nat. Med.</u> , 1997, 4:11-14, 16-17

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(37 CFR §1.98(b))			

Other Documents (include Author, Title, Date, and Place of Publication)		
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	AOO	Nitsch et al., "The Selective Muscarinic M1 Agonist AF102B Decreases Levels of Total A $\beta$ in Cerebrospinal Fluid of Patients with Alzheimer's Disease," <i>Ann. Neurol.</i> , 2000, 48:913-918
	APP	Ohyagi et al., "Selective increase in cellular A $\beta$ 42 is related to apoptosis but not necrosis," <i>NeuroReport.</i> , 2000, 11:167-171
	AQQ	Pagansonline.com, Chemical and Pharmaceutical Bulletin at <a href="http://www.pagansonline.com/pagansonline/cimicifuga.htm">http://www.pagansonline.com/pagansonline/cimicifuga.htm</a> .
	ARR	Petanceska and Gandy, "The Phosphatidylinositol 3-Kinase Inhibitor Wortmannin Alters the Metabolism of the Alzheimer's Amyloid Precursor Protein," <i>J. Neurochem.</i> , 1999, 73:2316-2320
	ASS	Pharmaceutical Information Network, "Black Cohosh," at <a href="http://www.pharminfo.com/cam/BlackCohoshch.html">http://www.pharminfo.com/cam/BlackCohoshch.html</a> .
	ATT	Querfurth and Selkoe, "Calcium Ionophore Increases Amyloid $\beta$ Peptide Production by Cultured Cells," <i>Biochem.</i> , 1994, 33(15):4550-4561
	AUU	Querfurth et al., "Caffeine Stimulates Amyloid $\beta$ -Peptide Release from $\beta$ -Amyloid Precursor Protein-Transfected HEK293 Cells," <i>J. Neurochem.</i> , 1997, 69:1580-1591
	AVV	Racchi and Govoni, "Rationalizing a pharmacological intervention on the amyloid precursor protein metabolism," <i>Trends Pharmacol. Sci.</i> , 1999, 20:418-423
	AWW	Sambamurti et al., "Glycosylphosphatidylinositol-anchored Proteins Play an Important Role in the Biogenesis of the Alzheimer's Amyloid $\beta$ -Protein," <i>J. Biol. Chem.</i> , 1999, 274(38):26810-26814
	AXX	Savage et al., "Turnover of Amyloid $\beta$ -Protein in Mouse Brain and Acute Reduction of Its Level by Phorbol Ester," <i>J. Neurosci.</i> , 1998, 18(5):1743-1752
	AYY	Shoji et al., "Production of the Alzheimer Amyloid $\beta$ Protein by Normal Proteolytic Processing," <i>Science</i> , 1992, 258:126-129
	AZZ	Suzuki et al., "An Increased Percentage of Long Amyloid $\beta$ Protein Secreted by Familial Amyloid $\beta$ Protein Precursor ( $\beta$ APP <sub>717</sub> ) Mutants," <i>Science</i> , 1994, 264:1336-1340
	AAAA	Viable Herbal Solutions, "Black Cohosh," at <a href="http://www.viable-herbal.com/herbdesc/1bcohash.ttm">http://www.viable-herbal.com/herbdesc/1bcohash.ttm</a> .
	ABBB	WebMD, Herbal Index, "Black Cohosh Root," at <a href="http://www.onhealth.webmd.com/alternatuive/resource/herbs/item,76985">http://www.onhealth.webmd.com/alternatuive/resource/herbs/item,76985</a> .
	ACCC	Whole Health Discount Center, "Black Cohosh," at <a href="http://www.health-pages.com/bc/index.html">http://www.health-pages.com/bc/index.html</a> .
	ADDD	Williams Nursury, "Cimicifuga," at <a href="http://www.Williams-nursury.com/cimicifuga.htm">http://www.Williams-nursury.com/cimicifuga.htm</a> .
	AEEE	Xu et al., "Estrogen reduces neuronal generation of Alzheimer $\beta$ -amyloid peptides," <i>Nat. Med.</i> , 1998, 4(4):447-451
	AFFF	Yager et al., "Natural Product Extracts that Reduce Accumulation of the Alzheimer's Amyloid $\beta$ Peptide," <i>Journal of Molecular Neuroscience</i> , 2002, 19:129-133.

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